PATENT COOPERATION TREATY

PCT

INTERNATIONAL SEARCH REPORT

(PCT Article 18 and Rules 43 and 44)

Applicant's or agent's file reference 3P/G-33319A/SZB	FOR FURTHER ACTION as	see Form PCT/ISA/220 ell as, where applicable, item 5 below.		
nternational application No.	International filing date (day/month/year)	(Earliest) Priority Date (day/month/year)		
PCT/EP2004/009321	19/08/2004	20/08/2003		
pplicant				
INTOG NO				
SANDOZ AG				
This International Search Report has according to Article 18. A copy is being	peen prepared by this International Searching g transmitted to the International Bureau.	Authority and is transmitted to the applicant		
	ists of a total of sheets.			
X It is also accompanied	by a copy of each prior art document cited in	this report.		
Basis of the report				
 With regard to the language, flanguage in which it was filed, 	the international search was carried out on the unless otherwise indicated under this item.	basis of the international application in the		
The internation this Authority (nal search was carried out on the basis of a tra	anslation of the international application furnished to		
		sed in the international application, see Box No. I.		
		in the international application, see Box 140. 1.		
2. Certain claims were f	ound unsearchable (See Box II).			
3. Unity of invention is	lacking (see Box III).			
4. With regard to the title,				
X the text is approved as	submitted by the applicant.			
the text has been estal	blished by this Authority to read as follows:			
. With regard to the abstract,				
	submitted by the applicant.			
the text has been estat	olished, according to Rule 38.2(b), by this Auth	nority as it appears in Box No. IV. The applicant		
may, within one month	from the date of mailing of this international se	earch report, submit comments to this Authority.		
. With regard to the drawings,				
	e published with the abstract is Figure No	-		
	y the applicant. this Authority, because the applicant failed to	Suggest a figure		
	this Authority, because this figure better chara	- · ·		
	be published with the abstract.			

International Application No PCT/EP2004/009321

PCT/EP2004/009321 A. CLASSIFICATION OF SUBJECT MATTER IPC 7 C12P21/02 C12P21/02 According to International Patent Classification (IPC) or to both national classification and IPC **B. FIELDS SEARCHED** Minimum documentation searched (classification system followed by classification symbols) C12P C12N IPC 7 C07K Documentation searched other than minimum documentation to the extent that such documents are included in the fields searched Electronic data base consulted during the international search (name of data base and, where practical, search terms used) EPO-Internal, WPI Data, PAJ, BIOSIS, FSTA C. DOCUMENTS CONSIDERED TO BE RELEVANT Category ° Citation of document, with indication, where appropriate, of the relevant passages Relevant to claim No. Χ HART R A ET AL: "Large scale, in situ 1-3,6,9, isolation of periplasmic IGF-I from E. 16-21.23 coli" BIO/TECHNOLOGY, vol. 12, November 1994 (1994-11), pages 1113-1117, XP001153305 ISSN: 0733-222X abstract page 1116, right-hand column, lines 3-50 WO 01/94585 A (CHAPMAN ANDREW PAUL; KING Α 1 DAVID JOHN (GB); ATHWAL DILJEET SINGH (GB);) 13 December 2001 (2001-12-13) cited in the application page 35, line 31 - page 36, line 8 page 34, line 29 - page 35, line 17 page 38, lines 4-31 page 40, lines 7-29 Further documents are listed in the continuation of box C. Patent family members are listed in annex. Special categories of cited documents: "T" later document published after the international filing date or priority date and not in conflict with the application but cited to understand the principle or theory underlying the "A" document defining the general state of the art which is not considered to be of particular relevance invention earlier document but published on or after the international "X" document of particular relevance; the claimed invention cannot be considered novel or cannot be considered to filing date document which may throw doubts on priority claim(s) or which is cited to establish the publication date of another citation or other special reason (as specified) involve an inventive step when the document is taken alone "Y" document of particular relevance; the claimed invention cannot be considered to involve an inventive step when the document is combined with one or more other such documents, such combination being obvious to a person skilled document referring to an oral disclosure, use, exhibition or other means document published prior to the international filing date but later than the priority date claimed in the art "&" document member of the same patent family Date of the actual completion of the international search Date of mailing of the international search report 16 December 2004 10/03/2005 Name and mailing address of the ISA Authorized officer European Patent Office, P.B. 5818 Patentlaan 2 NL – 2280 HV Rijswijk Tel. (+31–70) 340–2040, Tx. 31 651 epo nl,

Form PCT/ISA/210 (second sheet) (January 2004)

Fax: (+31-70) 340-3016

3

van de Kamp, M

International Application No
PCT/EP2004/009321

C.(Continu	ation) DOCUMENTS CONSIDERED TO BE RELEVANT	PC1/EP2004/009321
Category °		Relevant to claim No.
A	WO 93/06217 A (GENENTECH INC) 1 April 1993 (1993-04-01) page 30, line 30 - page 31, line 15	1
Т	ANDERSEN D C ET AL: "Production technologies for monoclonal antibodies and their fragments" CURRENT OPINION IN BIOTECHNOLOGY, vol. 15, no. 5, October 2004 (2004-10), pages 456-462, XP004588033 ISSN: 0958-1669 page 456, right-hand column, line 3 - page 457, left-hand column, line 12	
X	EP 0 177 343 A (GENENTECH INC) 9 April 1986 (1986-04-09) page 6, line 33 - page 7, line 12	1,6, 9-12, 16-23
	page 7, line 20 - page 9, line 6 page 20, line 15 - page 22, line 4 example 8 claims 12,13,15,16	
X	WO 03/004599 A (PANCER ZEEV; PELEG YOAV (IL); INSIGHT STRATEGY & MARKETING L (IL)) 16 January 2003 (2003-01-16) examples 2,3	1-5,14, 15,17-23
A	EP 0 974 654 A (MITSUI CHEMICALS INC) 26 January 2000 (2000-01-26) paragraphs '0066!, '0067! examples 1-3	1
١	EP 0 587 427 A (MITSUI TOATSU CHEMICALS) 16 March 1994 (1994-03-16) examples 3,4	1
4	DE OLIVEIRA J E ET AL: "High-yield purification of biosynthetic human growth hormone secreted in Escherichia coli periplasmic space" JOURNAL OF CHROMATOGRAPHY A, ELSEVIER SCIENCE, NL, vol. 852, no. 2, 13 August 1999 (1999-08-13), pages 441-450, XP004177405 ISSN: 0021-9673 paragraph 2.2	
	WO 01/57217 A (CHOI KI DOO; KWON SE CHANG (KR); BAE SUNG MIN (KR); JUNG SUNG YOUB (K) 9 August 2001 (2001-08-09) comparative example 2 examples 3,4 table I	1
	 -/	

3

International Application No
PCT/EP2004/009321

0./0	DOCUMENTO CONCUENTS TO BE OF FIVE	<u> </u>
Category °	ation) DOCUMENTS CONSIDERED TO BE RELEVANT Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
А	HANNIG G ET AL: "Strategies for optimizing heterologous protein expression in Escherichia coli" TRENDS IN BIOTECHNOLOGY, vol. 16, no. 2, February 1998 (1998-02), pages 54-60, XP004107042 ISSN: 0167-7799 figure 2 page 58, left-hand column, lines 13-45	
A	MIDDELBERG A P J: "Process-scale disruption of microorganisms" BIOTECHNOLOGY ADVANCES, vol. 13, no. 3, 1995, pages 491-551, XP004045404 ISSN: 0734-9750 the whole document	
A	BANEYX F: "Recombinant protein expression in Escherichia coli" CURRENT OPINION IN BIOTECHNOLOGY, vol. 10, no. 5, October 1999 (1999-10), pages 411-421, XP001041299 ISSN: 0958-1669 page 418, left-hand column, lines 4-49	

3

Information on patent family members

International Application No
PCT/EP2004/009321

Patent document cited in search report		Publication date		Patent family member(s)	-	Publication date
WO 0194585		13-12-2001	AU	6051101	A	17-12-2001
	••	10 12 2001	BG	106278		29-12-2002
			BR	0106682		14-05-2002
			CA	2380298		13-12-2001
			CN	1383450		04-12-2002
			CZ	20020837		15-05-2002
			DE	10192353		22-05-2003
			EP	1287140		05-03-2003
			WO	0194585		13-12-2001
			GB	2366800		20-03-2002
			HU	0202346		28-10-2002
			JP	2003535591		02-12-2003
				PA01013440		
			MX			04-09-2003
			NO	20020554		08-04-2002
			NZ	516596		30-07-2004
			PL	353960		15-12-2003
			SK	3152002		02-07-2002
			US	2003026805		06-02-2003
			US	2002151682		17-10-2002
			ZA	200200097 	A 	06-01-2003
WO 9306217	Α	01-04-1993	CA	2116774		01-04-1993
			EP	0604580	A1	06-07-1994
			EP	0861893	A2	02-09-1998
			JP	6510904	T	08-12-1994
			JP	2004041240	Α	12-02-2004
			WO	9306217	A1	01-04-1993
			US	5648237	Α	15-07-1997
EP 0177343	A	09-04-1986	US	4680262	Α	14-07-1987
			US	4963495	A	16-10-1990
			ĀŤ	78515		15-08-1992
			DE	3586386		27-08-1992
			DE	3586386		14-01-1993
			ĒΡ	0177343		09-04-1986
			JР	2521413		07-08-1996
			JP	6296491		25-10-1994
			JP	2102975		22-10-1994
			JP	8015440		21-02-1996
			JP	61092575		10-05-1986
			JP	2575642		29-01-1997
			JP	62500554		05-03-1987
			KR	9402814		05-03-1987
WO 03004599	Α	16-01-2003	WO	03004599	A2	16-01-2003
EP 0974654	Α	26-01-2000	AU	731758	 В2	05-04-2001
			AU	3801899		03-02-2000
			EP	0974654		26-01-2000
			ĴΡ	2000078989		21-03-2000
			KR	2000078383		25-02-2000
			US	6160089		12-12-2000
			US	6436674		20-08-2002
	 А	16-03-1994	DE	69318909	 D1	09-07-1998
FP 0587427	~	10-03-1994				
EP 0587427	•		חר	£0210000		
EP 0587427	,,		DE	69318909		24-12-1998
EP 0587427			DE EP JP	69318909 0587427 3398427	A1	16-03-1994 21-04-2003

Information on patent family members

International Application No
PCT/EP2004/009321

Patent document cited in search repor	ı	Publication date		Patent family member(s)	Publication date
EP 0587427	Α		JP	6269292 A	27-09-1994
			US	5496713 A	05-03-1996
WO 0157217	Α	09-08-2001	KR	2001073658 A	01-08-2001
			ΑU	762053 B2	19-06-2003
			AU	2890901 A	14-08-2001
			BR	0107708 A	19-11-2002
			CA	2397527 A1	09-08-2001
			CN	1418251 T	14-05-2003
			EP	1250439 A1	23-10-2002
			JP	2003521925 T	22-07-2003
			WO	0157217 A1	09-08-2001
			NZ	520076 A	25-07-2003
			RU	2230116 C2	10-06-2004
			ÜS	2004151695 A1	05-08-2004